REMARKS

The Rejections under 35 USC 112, second paragraph

Claim 1 is amended to recite the specific heteroaryls from claim 16 at each occurrence. Claim 16 is simultaneously cancelled. Applicants however disagree with the Examiner for reasons of record. One of skill in the art based on the disclosure would not find the "heteroaryl" term indefinite.

The allegedly indefinite term "contains" appearing at several places in the context of listing heteroatoms in claim 1 is replaced by "has."

"B" is changed to "Y" in order to prevent the possible alleged confusion regarding "B" being possibly understood to mean boron.

Applicants bring to the attention of the Examiner to a mistake in now cancelled claim 16. Claim 16 erroneously listed species of both heteroaryls and saturated heterocycles under the common name of heteroaryls. For a list for each group in the specification please see page 8. The currently amended claims do not contain this mistake.

Furthermore, during the previous reply, claim 1 was by mistake amended to narrow the definitions of R^2 to R^5 to H and of R^6 to methyl. The definitions are restored. However in view of the section 112 rejections to other parts of the claim, the term heteroaryl is amended as at other locations in the claim as discussed above. The present scope of the claims has been Examined as evidenced by the Office Action of February 19, 2002. No new searching burden is placed on the Examiner by the restoration of the claim to its proper scope. Applicant are very sorry for any inconvenience this may cause the Examiner.

The rejections over claim 6 should be overcome by the restoration of the definitions of R^1 and R^2 together. The $-(CHR^9)_n-NR^7-A-NR^8-Y$ group can attach to any member of the ring.

The Rejections under 35 USC 112, first paragraph

The section 112, first paragraph, rejections are overcome by the amendment to claim 1 by reciting the specific heteroaryls from claim 16 at each occurrence. Applicants however disagree with the Examiner for reasons of record. One of skill in the art based on the disclosure would have understood the scope of the claim and would have been fully able to practice the invention without undue experimentation.

Information Disclosure Statement

Applicants thank the Examiner for bringing the proviso to applicant's attention. Upon review, it appears that the source of the proviso, US 6,365,736, is not of record in the present application. An IDS is filed.

Attached hereto is a marked-up version of the changes made to the claims by the current amendment. The attached page is captioned **"Version With Markings To Show Changes Made"**.

The Commissioner is hereby authorized to charge any fees associated with this response or credit any overpayment to Deposit Account No. 13-3402.

Respectfully submitted,

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Date: October 17, 2002
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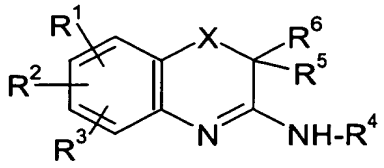
Version With Markings To Show Changes Made

In the Claims

The claims have been amended as follows:

1. (Twice Amended) A compound of Formula I, or a tautomeric or isomeric form or a salt of a compound of Formula I,



(I)

wherein

X is O,

R^1 is ~~$-(CHR^9)_n-NR^7-A-NR^8-B$~~ $-(CHR^9)_n-NR^7-A-NR^8-Y$,

~~R^2, R^3, R^4 and R^5 are hydrogen,~~

~~R^6 is methyl,~~

R^2 is hydrogen, or

R^1 and R^2 together with two adjacent carbon atoms form a 5-, 6-, 7- or 8-membered ring, which is monocyclic or bicyclic, saturated or unsaturated and in which 1 or 2 CH_2 groups can be replaced by oxygen or carbonyl, and which is substituted with $(CHR^9)_r-NR^7-A-NR^8-Y$, and is optionally substituted with C_{1-4} alkyl,

R^3 is hydrogen, halogen, NO_2, cyano, CF_3, $-OCF_3$, $-S-R^9$, $-O-R^9$, C_{3-7} cycloalkyl, $-NR^9-C(=NR^{10})-R^{11}$, $-NH-CS-NR^{12}R^{13}$, $-NH-CO-NR^{12}R^{13}$, $-CO-R^{14}$, $NR^{15}R^{16}$, C_{6-10} aryl, which optionally is substituted with halogen, cyano, C_{1-4} alkyl, $-S-R^9$, or $-O-R^9$, or is thienyl, imidazole, indole, isooxazole, isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene, thiadiazole, benzimidazole, benzofuran, benzoxazole, isoquinoline, quinoline, 2-C_{1-6} alkyl-3-amino-1,4-benzoxazine, or is 2-C_{1-6}-alkyl-3-keto-1,4-benzoxazine, or a C_{1-6} alkyl, which is optionally substituted with halogen, $-OR^9$, $-SR^9$, $-NR^{12}R^{13}$, $=NR^{12}$, $=NOC_{1-6}$ alkyl, =N-NHaryl, phenyl, C_{3-7} cycloallkyl or with thienyl, imidazole, indole, isooxazole,

isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene, thiadiazole, benzimidazole, benzofuran, benzoxazole, isoquinoline, quinoline, or is a C_{2-6} alkinyl, which is optionally substituted with halogen, $CONH_2$, C≡N or phenyl,

R^4 is hydrogen or acyl,

R^5 and R^6, independently of one another, are hydrogen, C_{3-7} cycloalkyl, phenyl, C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl radicals, which are optionally and independently of one another substituted with halogen, OH, O-C_{1-6} alkyl, SH, S-C_{1-6} alkyl, $NR^{15}R^{16}$, thienyl, imidazole, indole, isooxazole, isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene, thiadiazole, benzimidazole, benzofuran, benzoxazole, isoquinoline, quinoline, phenyl or C_{3-7} cycloalkyl,

R^7 is hydrogen, C_{1-6} alkyl, which is optionally substituted with phenyl, $COOC_{1-6}$ alkyl or CO-C_{1-6} alkyl,

R^8 is hydrogen, C_{1-6} alkyl, which is optionally substituted with phenyl, $COOC_{1-6}$ alkyl or COC_{1-6} alkyl,

A is a straight-chain or branched C_{1-6} alkylene, straight-chain or branched C_{1-6} alkenylene or $-(CH_2)_p$-Q-$(CH_2)_q$-,

~~B~~ Y is hydrogen or $-(CH_2)_p$-U,

Q is C_{3-7} cycloalkyl, indanyl, 5-, 6- or 7-membered saturated heterocycloalkyl with 1-2 N, O or S atoms, C_6-C_{10} aryl or ~~5- or 6-membered heteroaryl with 1-3 N, O or S atoms, which is optionally anellated with benzene~~ thienyl, imidazole, indole, isooxazole, isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene, thiadiazole, benzimidazole, benzofuran, benzoxazole, isoquinoline, quinoline, 2-C_{1-6} alkyl-3-amino-1,4-benzoxazine, or 2-C_{1-6}-alkyl-3-keto-1,4-benzoxazine,

U is hydrogen, C_{1-6} alkyl optionally substituted with halogen, C_{3-7} cycloalkyl, indanyl, C_{7-10} bicycloalkyl, C_{6-10} aryl or ~~5- or 6-membered heteroaryl with 1-3 N, O or S atoms, which is optionally anellated with benzene~~ thienyl, imidazole, indole, isooxazole, isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene,

thiadiazole, benzimidazole, benzofuran, benzoxazole, isoquinoline, quinoline, 2-C_{1-6} alkyl-3-amino-1,4-benzoxazine, or 2-C_{1-6}-alkyl-3-keto-1,4-benzoxazine, wherein the aryl or ~~heteroaryl radical~~ thienyl, imidazole, indole, isooxazole, isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene, thiadiazole, benzimidazole, benzofuran, benzoxazole, isoquinoline, quinoline, 2-C_{1-6} alkyl-3-amino-1,4-benzoxazine, or 2-C_{1-6}-alkyl-3-keto-1,4-benzoxazine, is optionally substituted with halogen, C_{1-4} alkyl, C_{1-4} alkoxy, CF_3, NO_2, NH_2, $N(C_{1-4}$ alkyl$)_2$, cyano, $CONH_2$, -O-CH_2-O-, -O-$(CH_2)_2$-O-, SO_2NH_2, OH, phenoxy or $COOC_{1-4}$ alkyl,

R^8 and ~~B~~ Y together with the nitrogen atom optionally form a 5- to 7-membered saturated heterocycle, which optionally ~~contains~~ has another oxygen, nitrogen or sulfur atom and is optionally substituted with C_{1-4} alkyl, phenyl, benzyl or benzoyl or form an unsaturated 5-membered heterocycle, which optionally ~~contains~~ has 1-3 N atoms and is optionally substituted with phenyl, C_{1-4} alkyl or halogen,

R^7 and A together with the nitrogen atom optionally form a 5- to 7-membered saturated heterocycle, which optionally ~~contains~~ has another oxygen, nitrogen or sulfur atom or ~~forms~~ form an unsaturated 5-membered heterocycle, which optionally ~~contains~~ has 1-3 N atoms,

m is 0, 1 or 2,

n and r is 0, 1 to 6,

p and q is 0 to 6,

R^9 and R^{10} is hydrogen or C_{1-6} alkyl,

R^{11} is C_{1-6} alkyl, $-NH_2$, -NH-CH_3, -NH-CN, C_{6-10} aryl optionally substituted with halogen, C_{1-4} alkyl or CF_3, or ~~5- or 6-membered heteroaryl with 1 to 4 nitrogen, sulfur or oxygen atoms that is optionally~~ an unsubstituted or substituted with halogen, C_{1-4} alkyl or CF_3, group selected from the group consisting of thienyl, imidazole, indole, isooxazole, isothiazole, furan, oxadiazole, oxazole, pyrazine, pyridazine, pyrimidine, pyridine, pyrazole, pyrrole, tetrazole, thiazole, triazole, thiophene, thiadiazole, benzimidazole, benzofuran, benzoxazole,

isoquinoline, quinoline, 2-C_{1-6} alkyl-3-amino-1,4-benzoxazine, and 2-C_{1-6}-alkyl-3-keto-1,4-benzoxazine,

R^{12} and R^{13} are hydrogen, C_{1-6}, alkyl, phenyl optionally substituted with halogen or C_{1-4} alkyl, benzyl optionally substituted with halogen or C_{1-4} alkyl, or C_{3-7} cycloalkyl,

R^{14} is hydrogen, hydroxy, C_{1-6} alkoxy, phenyl, C_{1-6} alkyl optionally substituted with CO_2H, CO_2C_{1-6} alkyl, hydroxy, C_{1-4} alkoxy, halogen, $NR^{15}R^{16}$, $CONR^{12}R^{13}$, phenyl, or C_{2-6} alkenyl optionally substituted with phenyl, cyano, $CONR^{12}R^{13}$ or CO_2C_{1-4} alkyl,

R^{15} and R^{16} are hydrogen, C_{1-6} alkyl, phenyl or benzyl, and

R^{15} and R^{16} together with the nitrogen atom optionally form a saturated 5-, 6-, or 7-membered ring, which optionally ~~contains~~ has another nitrogen, oxygen or sulfur atom and is optionally substituted with C_{1-4} alkyl, phenyl, benzyl or benzoyl,

wherein when R^6 is methyl and R^2, R^3, R^4 and R^5 are hydrogen, R^1 is not 6-((4-aminobenzyl)aminomethyl), 6-((4-dimethylaminobenzyl)aminomethyl), 6-((4-aminobenzyl) (tert-butyloxycarbonyl)aminomethyl), or 6-((4-dimethylaminobenzyl) (tert-butyloxycarbonyl)aminomethyl).

6. (Three Times Amended) A compound according to claim 1, wherein R^1 and R^2 together with two adjacent carbon atoms form ~~a~~ the 3- to 8-membered ring, that is substituted with ~~$-(CHR^9)_r-NR^7-A-NR^8B$~~ $-(CHR^9)_r-NR^7-A-NR^8Y$.

Claims 19-22 have been newly added.